

# Journal Pre-proof



Low-profile Zenith Alpha™ Thoracic stent graft modification using pre-loaded wires for urgent repair of thoracoabdominal and pararenal abdominal aortic aneurysms

Sukgu M. Han, MD, Emanuel R. Tenorio, MD, PhD, Aleem Mirza, MD, Louis Zhang, MD, Salome Weiss, MD, Gustavo S. Oderich, MD

PII: S0890-5096(20)30222-3

DOI: <https://doi.org/10.1016/j.avsg.2020.02.022>

Reference: AVSG 4935

To appear in: *Annals of Vascular Surgery*

Received Date: 23 October 2019

Revised Date: 12 February 2020

Accepted Date: 17 February 2020

Please cite this article as: Han SM, Tenorio ER, Mirza A, Zhang L, Weiss S, Oderich GS, Low-profile Zenith Alpha™ Thoracic stent graft modification using pre-loaded wires for urgent repair of thoracoabdominal and pararenal abdominal aortic aneurysms, *Annals of Vascular Surgery* (2020), doi: <https://doi.org/10.1016/j.avsg.2020.02.022>.

This is a PDF file of an article that has undergone enhancements after acceptance, such as the addition of a cover page and metadata, and formatting for readability, but it is not yet the definitive version of record. This version will undergo additional copyediting, typesetting and review before it is published in its final form, but we are providing this version to give early visibility of the article. Please note that, during the production process, errors may be discovered which could affect the content, and all legal disclaimers that apply to the journal pertain.

© 2020 Elsevier Inc. All rights reserved.

1 **Low-profile Zenith Alpha™ Thoracic stent graft modification using pre-loaded wires**  
2 **for urgent repair of thoracoabdominal and pararenal abdominal aortic aneurysms**

3

4 Sukgu M. Han, MD<sup>1\*</sup>, Emanuel R. Tenorio, MD, PhD<sup>2\*</sup>, Aleem Mirza, MD<sup>2</sup>, Louis

5 Zhang, MD<sup>1</sup>, Salome Weiss, MD<sup>2,3</sup> and Gustavo S. Oderich, MD<sup>2</sup>

6 *\* Shared first authorship*

7

8 <sup>1</sup>1520 San Pablo St, Suite 4300, Division of Vascular Surgery and Endovascular Therapy,

9 University of Southern California, Los Angeles, CA 90033;

10

11 <sup>2</sup>Gonda 4 South 200 First Street SW, Division of Vascular and Endovascular Surgery,

12 Mayo Clinic, Rochester, MN 55905;

13

14 <sup>3</sup>Freiburgstrasse, Department of Cardiovascular Surgery, Inselspital, Bern University

15 Hospital, University of Bern, 3010 Bern, Switzerland

16

17 Address for correspondence: Gustavo S. Oderich MD; Gonda Vascular Center, Mayo

18 Clinic; 200 First Street SW, Rochester MN 55905; Phone: (507) 2841575; FAX: (507)

19 266-7156; e-mail: [oderich.gustavo@mayo.edu](mailto:oderich.gustavo@mayo.edu)

20

21

22

23

24 **Abstract**

25 **Purpose:** To describe a modification technique using the low-profile Cook Zenith  
26 Alpha™ thoracic stent graft, and addition of a preloaded wire system, for urgent repair of  
27 pararenal (PRA) and thoracoabdominal (TAAA) aortic aneurysms.

28 **Methods:** We analyzed 20 consecutive patients who underwent urgent physician  
29 modified endograft (PMEG) repair of PRA and TAAA at two institutions. The low-  
30 profile Cook Zenith Alpha™ Thoracic stent graft was modified in according with each  
31 specific patient anatomic characteristics. Endpoints were technical success, 30-day  
32 mortality and major adverse events (MAEs).

33 **Results:** Technical success was achieved in all patients (100%). A total of 76 renal-  
34 mesenteric arteries were incorporated by fenestrations (70%) or directional branches  
35 (30%) with an average of  $3.7 \pm 0.6$  vessels per patient. There were six different types of  
36 stent configuration. The most common design consisted of four fenestrations (nine  
37 patients, 45%). The average of modification time was  $110 \pm 27$  minutes. Total procedure  
38 time (including the time for open component) was  $242 \pm 75$  minutes. There was no death  
39 within the first 30-day or hospital stay. MAEs occurred in 10 patients (50%). The most  
40 common MAEs were acute kidney injury (by RIFLE criteria) in six patients (30%), EBL  
41  $>1$  L and respiratory failure requiring reintubation in two patients (10%) each, paraplegia  
42 and ischemic colitis in one patient (5%) each. One patient (5%) required temporary, new-  
43 onset dialysis.

44 **Conclusion:** PMEG using low – profile Zenith Alpha™ thoracic stent graft was safe with  
45 no early mortality and acceptable early morbidity.

46

47 **Keywords:** Fenestrated Endovascular Aortic Aneurysm Repair; Physician Modified  
48 Endografts; Thoracoabdominal Aortic Aneurysm; Low Profile; Preloaded wires.

49

50

51

## 52 **Introduction**

53 Physician modified endograft (PMEG) technique provides a means for urgent  
54 total endovascular repair of pararenal (PRA) or thoracoabdominal aortic aneurysms  
55 (TAAAs) in patients who cannot wait for manufactured fenestrated and branched  
56 endografts. Previously reported PMEG techniques utilize the Zenith TX2<sup>®</sup> thoracic stent  
57 graft (Cook Medical Inc, Bloomington, Ind) as the base graft.<sup>1-4</sup> Further refinements of  
58 the modification technique include addition of diameter reducing wire<sup>5</sup>, mini-cuffs<sup>1</sup>,  
59 directional side branches and pre-loaded wires.<sup>2</sup> Introduced in the U.S. market in 2015,  
60 the low-profile Zenith Alpha<sup>™</sup> (16~20 French) thoracic stent graft (Cook Medical Inc,  
61 Bloomington, Ind) has replaced the Zenith TX2<sup>®</sup> (20~24 French) as the main workhorse  
62 off-the-shelf thoracic stent graft at most centers.<sup>6,7</sup> We describe our most recent PMEG  
63 technique using the low-profile Zenith Alpha<sup>™</sup> thoracic stent graft, incorporating the pre-  
64 loaded wire system, for urgent repair of PRAs and TAAAs.

65

## 66 **Methods**

### 67 *Patient Selection*

68 A retrospective two-center study was undertaken. The study was approved by the  
69 Institutional Review Board at each institution. Based on the retrospective nature of the

70 study, individual patient consent for the study was waived. Both institutions have  
71 physician-sponsored investigational device exemption (PS-IDE) studies, but in one  
72 institution eight cases were performed outside the PS-IDE study and under  
73 compassionate use due to inability to meet the inclusion and exclusion criteria. Off-label  
74 nature of the procedure, as well as other treatment options were disclosed to the patients  
75 during procedural consent. From March 2016 to January 2019, 20 patients underwent  
76 urgent PMEG repair of PRA and TAAAs at two institutions. Patients with high risk for  
77 open repair and presented with large, rapidly expanding, or symptomatic aneurysms, in  
78 which the requisite time for custom manufactured devices carried a significant risk of  
79 rupture were included in the study. PMEG repairs were not performed on patients with  
80 ruptured aneurysms presenting with hemodynamic instability. In these patients,  
81 induction of general anesthesia or graft modification time carried significant risk of total  
82 circulatory collapse.

### 83 ***Device design and planning***

84 Aneurysm morphology was determined by high-resolution computed tomography  
85 angiography (CTA) and devices designed with a minimum landing zone of 25mm. Three  
86 dimensional multi-planar as well as centerline reconstructions were performed for  
87 endograft modification planning. Endograft modifications were designed by surgeons  
88 with extensive experience in fenestrated and branched endografting, or dedicated training  
89 in complex aortic endografting. Patients with extent I-II TAAAs underwent staged repair  
90 with prophylactic spinal drainage. The first stage repair consisted of proximal thoracic  
91 stent graft placement, and the second stage fenestrated and branched endovascular repair  
92 using the PMEG technique. Neuromonitoring with motor evoked and sensory evoked

93 potential monitoring was indicated for patients with extent I-II TAAAs at the final  
94 procedure. Changes in the neuromonitoring triggers intraoperative maneuvers such as  
95 initiation (at 10 mmHg) or increase (down to 0-5 mmHg) of cerebrospinal fluid drainage  
96 and permissive hypertension with increments in target mean arterial pressure up to or  
97 above 90-100 mmHg. If these maneuvers do not reverse the changes in neuromonitoring,  
98 pelvic circulation is restored by retracting the femoral sheaths and allowing circulation to  
99 be restored to the hypogastric arteries and lower extremities. If neuromonitoring changes  
100 persist at the end of the procedure, temporary aneurysm sac perfusion can be allowed by  
101 leaving one of the visceral branches or the contralateral gate of the bifurcated device  
102 unstented.

103

#### 104 *Statistical analysis and definitions*

105 The reporting standards of the Society for Vascular Surgery were used to define  
106 stent-graft related outcomes and major adverse events (MAEs).<sup>(8,9)</sup> Major adverse events  
107 (MAEs) were defined using a composite end point, which included 30-day or in-hospital  
108 mortality, acute kidney injury (by RIFLE criteria), new-onset dialysis, myocardial  
109 infarction, respiratory failure requiring prolonged mechanical ventilation or reintubation,  
110 paraplegia, stroke, bowel ischemia requiring surgical resection or intensive medical care,  
111 and estimated blood loss > 1L. Outcome measurements were technical success, 30 – day  
112 mortality and MAEs. The results were reported as percent for categorical variables and  
113 mean  $\pm$  standard deviation (median, IQR) for continuous variables.

114

115

116 ***Technique***

117 To illustrate our technique, we present one case in detail. Patient 3 is a 77-year-  
118 old female who presented with a new onset abdominal and flank pain in the setting of a  
119 rapidly enlarging 7.5 cm Crawford extent II TAAA. The patient had a type 3 aortic arch,  
120 and moderate amount of aortic mural thrombus (**Figure 1**). The first stage operation  
121 consisted of a right iliofemoral bypass via a flank incision, and the placement of a tapered  
122 Zenith Alpha™ thoracic stent graft (38 ~ 34 x 167 mm). A prophylactic cerebrospinal  
123 drain was placed preoperatively.

124 ***Modification***

125 A tapered 36 ~ 32 x 209 mm proximal component Cook Zenith Alpha™ thoracic  
126 stent graft was modified using strict sterile technique (**Figure 2**). The stent graft was  
127 unsheathed and the proximal fixation barbs were removed (**Figure 2A-B**). Fenestrations  
128 were created at the predetermined locations using ophthalmic cautery (**Figure 2C**). Each  
129 fenestration was reinforced with double layers of Nitinol loops of an endovascular snare  
130 EnSNARE (Merit Medical, South Jordan, UT) using locking running 5-0 Ethibond  
131 (Ethicon, Cincinnati, OH) sutures (**Figure 2D**). Radiopaque gold markers (Accellent,  
132 Washington, Mass) were sutured around each fenestration (**Figure 2E**). The markers  
133 were also sutured at 12 and 6 o'clock of the stent graft to note the anterior and posterior  
134 position. Diameter reducing ties were added along the posterior wall of the stent graft  
135 (**Figure 2F**). All fenestrations were preloaded with 0.014-inch guidewires for the renal  
136 arteries or 0.018-inch guidewires for the mesenteric arteries (**Figure 2G**). The modified  
137 stent graft was resheathed using 2-0 Silk ties and Silastic tape (**Figure 2H-I, Video**).  
138 Side arm branches were created by sewing 1~2cm segment of 6mm (or 8mm self-

139 expanding Viabahn covered stents (W.L. Gore and Associates, Flagstaff, AZ) onto  
140 cauterized orifices, using running 6-0 Gore-Tex sutures (W.L. Gore and Associates,  
141 Flagstaff, AZ) (**Figure 3**).

142

### 143 *Implantation*

144 Under general anesthesia, left proximal brachial artery was surgically exposed  
145 through a 3cm longitudinal incision immediately distal to the axillary hairline. The right  
146 iliofemoral bypass graft was re-exposed in the groin. Percutaneous left femoral access  
147 was performed with preclose technique using Proglide devices (Abbott Vascular, Santa  
148 Clara, CA). The patient was fully heparinized. A 12Fr x 45cm Dryseal Flex<sup>®</sup> sheath  
149 (Gore and Associates, Flagstaff, AZ) was introduced from the left brachial approach and  
150 positioned in the descending thoracic aorta. Through-and-through brachiofemoral access  
151 was established, then a 5Fr x 90cm sheath was placed over this access. The preloaded  
152 wires were passed through the 5-Fr through-and-through sheath (**Figure 4A**). The  
153 modified Zenith Alpha<sup>™</sup> stent graft was introduced over the Lunderquist wire, and  
154 advanced to the paravisceral aorta with the 5-Fr sheath. The 5Fr sheath was removed  
155 from the brachial access. Under the guidance of Fusion CT (GE Discovery IGS 730, GE  
156 Healthcare, Chicago, IL), the stent graft was positioned with each fenestration in  
157 alignment with its target vessel (**Figure 4B**). The stent graft was partially deployed while  
158 individual visceral and renal branches were sequentially catheterized from the brachial  
159 approach, using the preloaded wires (**Figure 4C**). The branch catheterization technique  
160 using preloaded wires was previously described.<sup>(2)</sup> For the celiac axis (CA), 0.018-inch  
161 Steelcore wire was used, while 0.035-inch Amplatz and 0.035-inch Rosen wires were



162 used for the superior mesenteric artery (SMA), and renals. Following the complete target  
163 branch catheterization, the stent graft was fully deployed, and diameter-reducing wire  
164 removed. The renal arteries were stented with 5mm x 22mm iCAST covered stents  
165 (Atrium, Merrimack, NH) (**Figure 4D-E**). The CA and SMA were stented with 7mm x  
166 22mm and 7mm x 38mm iCAST, respectively (**Figure 4F-G**). All branch stents were  
167 flared to 10mm across the fenestrations. The repair was extended distally using 36mm  
168 Zenith aortic cuff (Cook Medical, Bloomington, IN). The completion angiography  
169 showed complete aneurysm exclusion with widely patent branch stents (**Figure 4H**).

## 170 **Results**

### 171 *Patient characteristics*

172 Mean age of our cohort was  $74\pm 8$  years (73, 68 –78), and 10 patients (50%) were  
173 male (**Table I**). Aneurysms were classified as TAAA in 16 patients (80%; Extent I in 1,  
174 II in 6, III in 5, and IV in 4) or PRA in 4 patients (20%). Mean aneurysm diameter was  
175  $74\pm 17$  mm (70, 63 – 88). The most prevalent cardiovascular risk factors were  
176 hypertension in 19 (95%) and prior history of cigarette smoking in 18 patients (90%).  
177 Ten patients (50%) presented with symptomatic non-ruptured or contained rupture of the  
178 aortic aneurysm.

### 179 *Stent-graft design*

180 There were six different types of stent configuration. The most common design,  
181 which was selected in 9 patients (45%), consisted of four fenestrations for the CA, SMA  
182 and renal arteries. Four patients had a single functioning kidney with chronic occlusion of  
183 one renal artery resulting in the incorporation of only three target vessels. A total of 74  
184 renal-mesenteric arteries were incorporated by fenestrations (70%) or directional

185 branches (30%) with an average of  $3.7\pm 0.6$  (4, 3 – 4) vessels per patient (**Table II**). All  
186 fenestrations and directional branches were loaded with 0.018 or 0.014-inch wires. The  
187 average modification time and length of the main device modified were  $110\pm 27$  (105, 96-  
188 131) minutes and  $179\pm 29$  (170, 161-207) millimeters, respectively.

### 189 *Procedure detail*

190 Technical success, defined by deployment of the aortic stent-graft and all intended  
191 side branch components, was achieved in 100% of patients (**Table II**). Twelve patients  
192 (60%) had prophylactic placement of cerebrospinal fluid drainage. Among eight patients  
193 that did not receive prophylactic spinal drain, aneurysm type was extent IV in four  
194 patients, pararenal in three and extent II in one. Mean endovascular procedure time was  
195  $166\pm 85$  minutes (118, 98 – 262) with fluoroscopy time of  $74\pm 27$  minutes (72, 59 – 89),  
196 and  $131\pm 52$  ml of iodinated contrast (120, 90 – 150) was used for each case. Total  
197 procedure time (including the time for open components) was  $242\pm 75$  minutes (215, 192  
198 – 281). Nine patients (45%) had the procedure in staged fashion with average of  $17\pm 29$   
199 (5, 2-21) days for the second stage, completion procedure.

### 200 *Early outcomes*

201 There was no death within the first 30-day or hospital stay. The mean length of  
202 hospital stay was  $9\pm 7$  days (7, 5 – 11). Ten patients (50%) had major adverse events  
203 (MAEs, **Table III**). The most common MAEs were acute kidney injury (by RIFLE  
204 criteria) in six patients (30%), EBL >1 L and respiratory failure requiring reintubation in  
205 two patients (10%) each, paraplegia and ischemic colitis in one patient (5%) each. One  
206 patient (5%) required temporary, new-onset dialysis. Two patients (10%) had a minor  
207 perioperative stroke (NIH Stroke Score 2). The patient with paraplegia, had four

208 directional branches endovascular repair of the symptomatic non-ruptured Extent II  
209 TAAA. He woke up neurologically intact from anesthesia. CSF drainage was open, and  
210 mean arterial pressure (MAP) was maintained > 80mmHg. Eight hours later, patient had  
211 an acute onset sharp back pain radiating to legs and became densely paraplegic. This  
212 never improved despite adjunctive measures.

213 Four patients required early secondary intervention to treat one type IA endoleak  
214 and three access complications. Follow-up imaging was obtained in all patients. After  
215 mean follow-up of  $5\pm 6$  (3, 1-9) months, there was no reintervention, aortic-related death  
216 or branch instability.

217

## 218 **Discussion**

219 Collective experience with fenestrated, branched stent graft technology is well  
220 established with implantation in thousands of patients, and continues to grow at a rapid  
221 pace. Concurrently, advances in low profile aortic stent graft technology have been  
222 incorporated to the manufactured fenestrated and branched devices. However,  
223 customized endografts take weeks for delivery and off-the-shelf devices for complex  
224 aortic anatomy are limited. Hence, the role of PMEGs continues to exist in urgent cases  
225 without viable open surgical options.

226 Reported outcomes by the authors regarding PMEGs have been favorable.  
227 Oderich and colleagues reported their series of 30 patients, 47% of which had TAAA.  
228 For a total of 85 fenestrations, branch stenting success was 98%, with one perioperative  
229 death. In this study, median follow-up was 14 months, with 1-year branch patency of  
230 97% and freedom from endoleak of 88%.<sup>(10)</sup> In 2012, Starnes and colleagues described

231 their initial outcomes of 47 consecutive patients with juxtarenal aneurysms treated with  
232 PMEG.<sup>11</sup> Technical success was 98%, with a total of 82 fenestrations, and a perioperative  
233 mortality of 2%. Since then, Starnes and colleagues have published their updated mid-  
234 term results now with 64 consecutive patients, 145 total fenestrations, and 4-year follow-  
235 up.<sup>12</sup> Technical success remained high at 95% and 30-day mortality of 5.1%.  
236 Complication rate was 12% including three myocardial infarctions, one stroke, one renal  
237 failure, and four respiratory failure. Freedom from stent-graft migration and freedom  
238 from rupture or conversion to open at 12 months was 100%. Only two patients had type I  
239 or type III endoleak at 12 months. No new endoleaks have been detected in patients seen  
240 at 4-year follow-up. Sac stability or shrinkage was seen in 97.7% of patients at one year  
241 and 95.2% of patients at three years. Mean follow up was 28 months, with four  
242 aneurysm-related deaths. All four aneurysm-related deaths were within the 30-day  
243 perioperative window, while an additional eight all-cause mortalities occurred during  
244 follow-up for an overall survival of 70% at 4 years.

245         Recent report from a high-volume aortic center in Hamburg, Germany  
246 demonstrated similarly high technical success of PMEG even in the setting of contained  
247 rupture.<sup>13</sup> Their cohort of 21 patients consisted of 11 TAAA, and 13 patients presenting  
248 with contained rupture. Technical success of PMEG implantation was achieved in 100%  
249 of the patients. Thirty-day survival was 95%. Two cases of permanent paralysis were in  
250 patients who had intraoperative hypotension due to rupture. There was 1 late aneurysm-  
251 related death in a patient who developed an aortoenteric fistula. At 11 months follow up,  
252 all branch stents remained patent. This report highlights that PMEG technique remains  
253 useful even in European centers where the off-the-shelf device (t-Branch) is available on

254 the market. The authors noted that “since customized endografts take weeks for delivery  
255 and off-the-shelf devices for complex aortic anatomy are limited, the surgeon-modified  
256 stent-grafts appears to be an effective endovascular option because it facilitates an  
257 anatomically correct reconstruction.”

258 O’Donnell and colleagues analyzed the outcomes of 1396 complex EVARs using  
259 the national Vascular Quality Initiatives database. Complex EVARs consisted of 880  
260 FEVARs using commercially available ZFEN (Cook Medical, Bloomington, IN), 256  
261 PMEGs, and 260 chimney/snorkel EVARS. Compared to ZFEN and chimney/snorkel  
262 EVARs, PMEGs were utilized to treat more extensive aneurysms, incorporating more  
263 branch vessels (mean:  $3.3\pm 0.8$  arteries vs.  $2.5\pm 0.6$  ZFEN, and  $1.9\pm 0.9$  chimney/snorkels)  
264 involving more celiac and SMA. However, PMEG had the lowest unadjusted  
265 perioperative death rate (2.7%), compared to ZFEN (3.4%) and chimney/snorkel (6.1%).  
266 Stroke rates after PMEG (0.9%) and ZFEN (0.8%) were similar, while chimney/snorkel  
267 EVARs had significantly higher stroke rates (3.3%,  $p=.03$ ). This study reflects that 1)  
268 PMEGs play a significant role in the real-world practice of complex EVARs in the  
269 United States, and 2) short-term results achieved by PMEG are similar to the excellent  
270 results achieved by ZFEN, and superior to snorkel/chimney EVARs.<sup>14</sup>

271 Since the published PMEG techniques using TX2 platform (Cook Medical,  
272 Bloomington, IN)<sup>1-4</sup>, the low-profile Zenith Alpha™ thoracic stent grafts have largely  
273 replaced the TX2 as the off-the-shelf thoracic stent graft at many centers. True urgent  
274 and emergent cases will most commonly involve use of readily available stent grafts. As  
275 such, familiarity with the low-profile Zenith Alpha™ thoracic stent graft and its

276 modification steps are critical components of achieving successful outcomes using  
277 PMEG technique.

278         There are several differences in the PMEG technique using the Zenith Alpha™  
279 thoracic stent graft, from its predecessor, TX2. First, widely spaced Nitinol stents on the  
280 Alpha device (**Figure 5**), compared to much tighter stainless-steel Z stents offer more  
281 space to place the fenestrations without having to bend and relocate the stent wires  
282 (**Figure 6**). Second, our approach to creating fenestrations involves efforts to reproduce  
283 the features of manufactured devices. Every fenestration is created to be 8x8mm or  
284 smaller and reinforced with a double layer of EnSNARE (Merit Medical, South Jordan,  
285 UT) wires constructed of braided Nitinol. The wires are secured around the fenestration  
286 by tightly spaced 5-0 Ethibond locking sutures (**Figure 2B**). These steps are aimed at  
287 maximizing durable seal across the fenestrations with flared balloon expandable covered  
288 stents. Sterile gold markers are placed in the configuration of the manufactured devices.  
289 Third, it is our preference to remove the active fixation barbs in order to facilitate  
290 intracorporeally rotation of the device.

291         Despite of the high-risk patients (100% of patients were American Society of  
292 Anesthesiology category III or IV) in the urgent or emergent setting, our results compare  
293 favorably with the pre-existing literature.<sup>12-16</sup> Technical success was achieved in 100%  
294 of patient with no 30-day mortality. Preloaded wire technique on the Zenith Alpha  
295 platform facilitates successful target vessel catheterization, as evidenced by mean  
296 fluoroscopy time of 74min, contrast amount of 131ml. This compares favorably to the  
297 largest PMEG experience previously published (fluoroscopy time: 108mins, and contrast:

298 191ml).<sup>15</sup> The rates of MAEs were acceptable with low secondary intervention. Branch  
299 patency was 100% after mean follow-up 5±6 months with no branch instability observed.

300 Despite these promising early results, some points of caution should be noted.  
301 Because there is no quality control and there is no oversight from industry representatives  
302 or a planning center, the technique is entirely dependent on the operator's experience  
303 with sizing, modifications, and device implantation. Modifications that are ill planned or  
304 poorly conducted can lead to disastrous complications and are often not reported in the  
305 literature. In addition, physician modifications of an approved device automatically  
306 exempt the manufacturer from any product liability claim. Although any physician can  
307 perform these modifications, the Food and Drug Administration (FDA) and the SVS do  
308 not endorse the technique unless it is performed under an investigational study. Last,  
309 PMEGs are costly and may not be fully reimbursed if they are performed outside these  
310 investigational studies.<sup>17</sup>

311 In conclusion, physician modified endograft technique can be applied to the low-  
312 profile Zenith Alpha™ thoracic stent graft, to enable urgent total endovascular repair of  
313 thoracoabdominal and complex aortic aneurysms in patients with high risk for open  
314 repair who cannot wait for manufactured devices.

315

316

317

318

319

320

321

322 **Reference**

323

- 324 1. Oderich GS, Fatima J, Gloviczki P. Stent graft modification with mini-cuff  
325 reinforced fenestrations for urgent repair of thoracoabdominal aortic aneurysms. *J Vasc*  
326 *Surg.* 2011;54(5):1522-6.
- 327 2. Oderich GS, Mendes BC, Correa MP. Preloaded guidewires to facilitate  
328 endovascular repair of thoracoabdominal aortic aneurysm using a physician-modified  
329 branched stent graft. *J Vasc Surg.* 2014;59(4):1168-73.
- 330 3. Sweet MP, Starnes BW, Tatum B. Endovascular treatment of thoracoabdominal  
331 aortic aneurysm using physician-modified endografts. *J Vasc Surg.* 2015;62(5):1160-7.
- 332 4. Scali ST, Neal D, Sollanek V, Martin T, Sablik J, Huber TS, et al. Outcomes of  
333 surgeon-modified fenestrated-branched endograft repair for acute aortic pathology. *J*  
334 *Vasc Surg.* 2015;62(5):1148-59 e2.
- 335 5. Oderich GS. Diameter-reducing wire to facilitate deployment of a modified  
336 Zenith fenestrated stent graft. *Ann Vasc Surg.* 2010;24(7):980-4.
- 337 6. Illig KA, Ohki T, Hughes GC, Kato M, Shimizu H, Patel HJ, et al. One-year  
338 outcomes from the international multicenter study of the Zenith Alpha Thoracic  
339 Endovascular Graft for thoracic endovascular repair. *J Vasc Surg.* 2015;62(6):1485-94  
340 e2.
- 341 7. Starnes BW, Dwivedi AJ, Giglia JS, Woo K, Yeh C, Investigators TS.  
342 Endovascular repair for blunt thoracic aortic injury using the Zenith Alpha low-profile  
343 device. *J Vasc Surg.* 2015;62(6):1495-503 e1.



- 344 8. Chaikof EL, Blankensteijn JD, Harris PL, White GH, Zarins CK, Bernhard VM,  
345 et al. Reporting standards for endovascular aortic aneurysm repair. *Journal of vascular*  
346 *surgery*. 2002;35(5):1048-60.
- 347 9. Fillinger MF, Greenberg RK, McKinsey JF, Chaikof EL. Reporting standards for  
348 thoracic endovascular aortic repair (TEVAR). *Journal of vascular surgery*.  
349 2010;52(4):1022-33. e5.
- 350 10. Oderich GS, Ricotta JJ. Modified fenestrated stent grafts: device design,  
351 modifications, implantation, and current applications. *Perspectives in vascular surgery*  
352 *and endovascular therapy*. 2009;21(3):157-67.
- 353 11. Starnes BW. Physician-modified endovascular grafts for the treatment of elective,  
354 symptomatic, or ruptured juxtarenal aortic aneurysms. *J Vasc Surg*. 2012;56(3):601-7.
- 355 12. Starnes BW, Heneghan RE, Tatum B. Midterm results from a physician-  
356 sponsored investigational device exemption clinical trial evaluating physician-modified  
357 endovascular grafts for the treatment of juxtarenal aortic aneurysms. *Journal of vascular*  
358 *surgery*. 2017;65(2):294-302.
- 359 13. Tsilimparis N, Heidemann F, Rohlfes F, Diener H, Wipper S, Debus ES, et al.  
360 Outcome of surgeon-modified fenestrated/branched stent-grafts for symptomatic complex  
361 aortic pathologies or contained rupture. *Journal of Endovascular Therapy*.  
362 2017;24(6):825-32.
- 363 14. O'donnell TF, Patel VI, Deery SE, Li C, Swerdlow NJ, Liang P, et al. The state of  
364 complex endovascular abdominal aortic aneurysm repairs in the Vascular Quality  
365 Initiative. *Journal of vascular surgery*. 2019.

- 366 15. Oderich GS, Ribeiro MS, Sandri GA, Tenorio ER, Hofer JM, Mendes BC, et al.  
367 Evolution from physician-modified to company-manufactured fenestrated-branched  
368 endografts to treat pararenal and thoracoabdominal aortic aneurysms. *Journal of vascular*  
369 *surgery*. 2018.
- 370 16. Ricotta II JJ, Tsilimparis N. Surgeon-modified fenestrated-branched stent grafts to  
371 treat emergently ruptured and symptomatic complex aortic aneurysms in high-risk  
372 patients. *Journal of vascular surgery*. 2012;56(6):1535-42.
- 373 17. Abel D, Farb A. Application of Investigational Device Exemptions regulations to  
374 endograft modification. *Journal of vascular surgery*. 2013;57(3):823-5.
- 375  
376  
377  
378  
379  
380  
381  
382  
383  
384  
385  
386  
387  
388  
389  
390  
391  
392  
393  
394  
395  
396  
397  
398  
399  
400  
401

402 **Table I.** Demographics, clinical and anatomical characteristics of 20 patients treated  
 403 using physician modified endograft (PMEG with Zenith Alpha™ thoracic stent graft).

<i>n</i> = number of patients	Overall
	<i>n</i> =20 (%) or mean ± SD (median, IQR)
<b>Demographics</b>	
Age (years old)	74±8 (73, 68-78)
Age > 80 years old	4 (20)
Male gender	10 (50)
<b>Cardiovascular risk factors</b>	
Cigarette smoking	18 (90)
Hypertension	19 (95)
Hypercholesterolemia	17 (85)
Coronary Artery Disease	11 (55)
COPD	10 (50)
Peripheral arterial disease	5 (25)
CKD Stage	
Stage I	2 (10)
Stage II	8 (40)
Stage III	8 (40)
Stage IV	2 (10)
CKD Stage III-IV	11 (55)
Diabetes Mellitus	1 (5)
Congestive Heart Failure	3 (15)
Stroke/TIA	3 (15)
Prior aortic repair	12 (60)
Ascending or arch	2 (10)
Open-Infrarenal	4 (20)
EVAR-AAA	1 (5)
TEVAR-1 Stg	9 (45)
<b>Preoperative evaluation</b>	
Serum Creatinine (mg/dl)	1.2±0.5 (1.0, 0.9-1.3)
eGFR (mL/min/1.73 m <sup>2</sup> )	61±22 (59, 49-81)
Body Mass Index (kg/m <sup>2</sup> )	27±7 (25, 22-30)
<b>Risk assessment</b>	
<b>ASA Score</b>	
Class 3	10 (50)
Class 4	10 (50)
<b>Anatomical Characteristics</b>	
Max aortic diameter	74±17 (70, 63-88)

Aneurysm Type	
Crawford extent I	1 (5)
Crawford extent II	6 (30)
Crawford extent III	5 (25)
Crawford extent IV	4 (20)
Pararenal	4 (20)
Post-dissection TAAA	1 (5)

---

404

405

406

407

408

409

410

411

412

413

414

415

416

417

418

419

420

421

**Table II.** Procedure detail and device design of 20 patients treated using physician modified

423 endograft (PMEG) with Zenith Alpha™ thoracic stent graft.

<i>n</i> = number patients	Overall
	<i>n</i> =20 (%) or mean ±SD (median, IQR)
General Anesthesia	20 (100)
Status of aneurysm	
Large/rapidly expanding aortic aneurysm	10 (50)
Symptomatic non-ruptured	9 (45)
Contained ruptured	1 (5)
Cerebrospinal Fluid Drainage	12 (60)
Somatosensory evoked potential/Motor evoked potentials	14 (70)
Preloaded wires	20 (100)
Reducing diameter tie	20 (100)
Brachial access	19 (95)
Left side	17 (85)
Right side	2 (10)
Percutaneous femoral approach	15 (75)
Unilateral	11 (55)
Bilateral	8 (40)
Iliac conduit	1 (5)
Modification time (min)	110±27 (105, 96-131)
Length of the main device modified (mm)	179±29 (170, 161-207)
Amount of contrast used (ml)	131±52 (120, 90-150)
Total endovascular time (min)	260±44 (271, 232-292)
Total fluoroscopy time (min)	74±27 (72, 59-89)
Total air kerma (mGy)	1975±1595 (1678, 707-3156)
Dose Area Product (Gy.cm <sup>2</sup> )	233±154 (158, 134-291)
Estimated blood loss (ml)	423±283 (350, 250-600)

Hospital stay (days)	9±7 (7, 5-11)
Technical success	20 (100)

---

<b>Device design</b>	<b>74</b>
Target vessel per patient	3.7±0.6 (4, 3-4)
Fenestration	52 (70)
Directional branches	22 (30)
Celiac axis	17 (23)
Fenestration	13 (18)
Directional branches	4 (5)
Superior mesenteric artery	20 (27)
Fenestration	14 (19)
Directional branches	6 (8)
Right renal artery	19 (26)
Fenestration	13 (18)
Directional branches	6 (8)
Left renal artery	17 (23)
Fenestration	11 (15)
Directional branches	6 (8)
Additional artery (Accessory RRA incorporated by fenestration)	1 (1)

---

424

425

426

427

428

429

430

431

**Table III.** Mortality and major adverse events (MAEs) < 30 days of 20 patients treated using physician modified endograft (PMEG) with Zenith Alpha™ thoracic stent graft.

	<b>Overall</b>
	<i>n=20 (percent)</i>
<i>n= number patients</i>	
<b>Early death</b>	0 (0)
<b>Any major adverse event</b>	10 (50)
Estimate blood loss higher than 1000 ml	2 (10)
Acute Kidney Injury by RIFLE criteria	6 (30)
New-Onset Dialysis	1 (5)
Myocardial Infarction	0 (0)
Respiratory failure	2 (10)
<b>Any spinal cord injury</b>	2 (10)
<i>Grading Classification</i>	
1-2 (paraparesis)	1 (5)
3a-c (paraplegia)	1 (5)
<i>Timing</i>	
Immediate	0 (0)
Delayed	2 (10)
<b>Any stroke</b>	2 (10)
Stroke major	0 (0)
Stroke minor/TIA	2 (10)
Bowel ischemia	1 (5)
Access complication	3 (15)

434 *RIFLE*, Risk, Injury, and Failure; and Loss; and End-stage kidney disease.

435

436

437

438

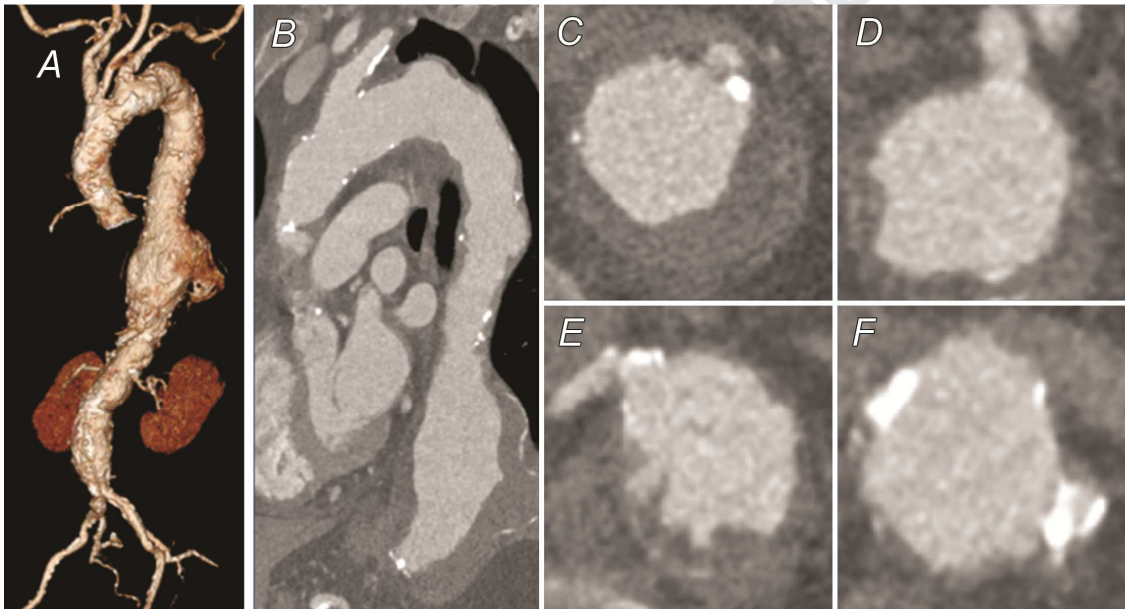
439

440

441

442

443 **Figure 1.** Computed tomography angiography demonstrated a 7.7cm extent II  
444 thoracoabdominal aortic aneurysm (**A**). The descending thoracic aorta had a moderate  
445 amount of mural thrombus (**B**). The celiac artery (**C**) and superior mesenteric artery (**D**)  
446 were patent. Increased amount of shaggy aortic mural thrombus was noted at the level of  
447 the right renal artery with moderate stenosis (**E**). The left renal artery had a high-grade  
448 stenosis with calcified ostial lesion (**F**). By permission of Mayo Foundation for Medical  
449 Education and Research. All rights reserved.



450

451

452

453

454

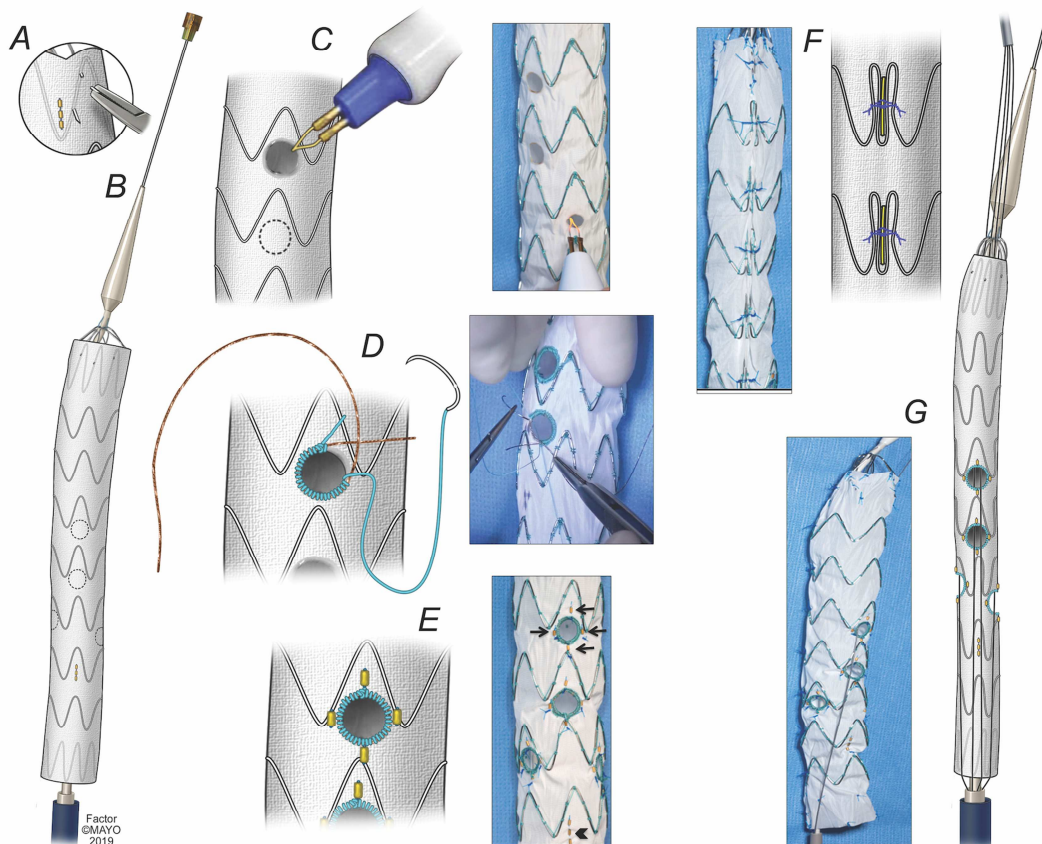
455

456

457



458 **Figure 2.** A tapered low profile alpha proximal component thoracic stent graft was  
 459 unsheathed. The device was modified on site by creating fenestrations for the visceral  
 460 and renal arteries (**A-C**). Fenestrations were reinforced with double layers of braided  
 461 nitinol wires using running 5-0 Ethibond sutures (Ethicon, Cincinnati, OH) (**D**). Each  
 462 fenestration was marked with four gold radiopaque markers (arrows). Longitudinal  
 463 anterior markers (arrow head) and transverse posterior markers were added (**E**).  
 464 Diameter reducing ties were added along the posterior aspect of the endograft (**F**). Each  
 465 fenestration was preloaded with a 0.014 wire (**G**). The device was constrained with Silk  
 466 ties and silastic tape, then reloaded into the sheath retrograde (**H-I**). By permission of  
 467 Mayo Foundation for Medical Education and Research. All rights reserved.





Factor  
©MAYO  
2019

469

470

471

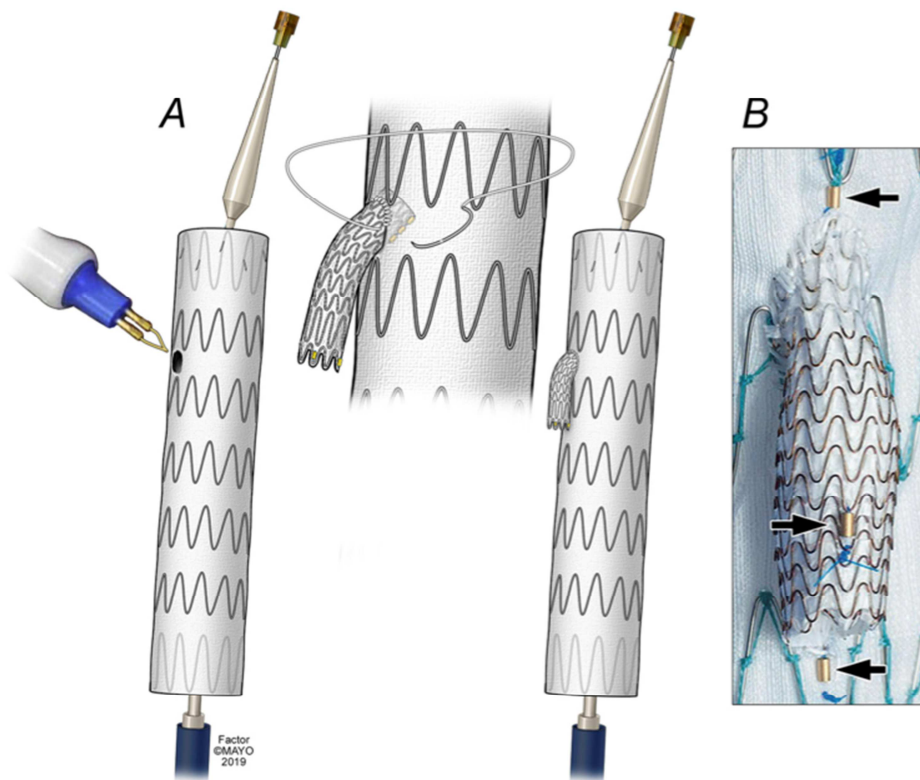
472

473

474

475

476 **Figure 3.** Directional branches are fashioned using 2-cm long Viabahn stent grafts (WL  
477 Gore, Flagstaff, AZ). The distal edge of the branch is positioned approximately 2 cm  
478 above the intended location of the target vessel within the same clock position. A  
479 Viabahn stent graft (**A**) is anastomosed end to side to the fenestration using running 5–0  
480 Gore sutures (WL Gore, Flagstaff, AZ). The distal edge of the Viabahn branch is stitched  
481 to the fabric of the TX2 stent graft to prevent movement during re-sheathing and  
482 deployment. Radiopaque markers are placed at the proximal and distal edge of the  
483 fenestration and at the distal edge of the branch (**B**, *black arrows*). By permission of  
484 Mayo Foundation for Medical Education and Research. All rights reserved.  
485



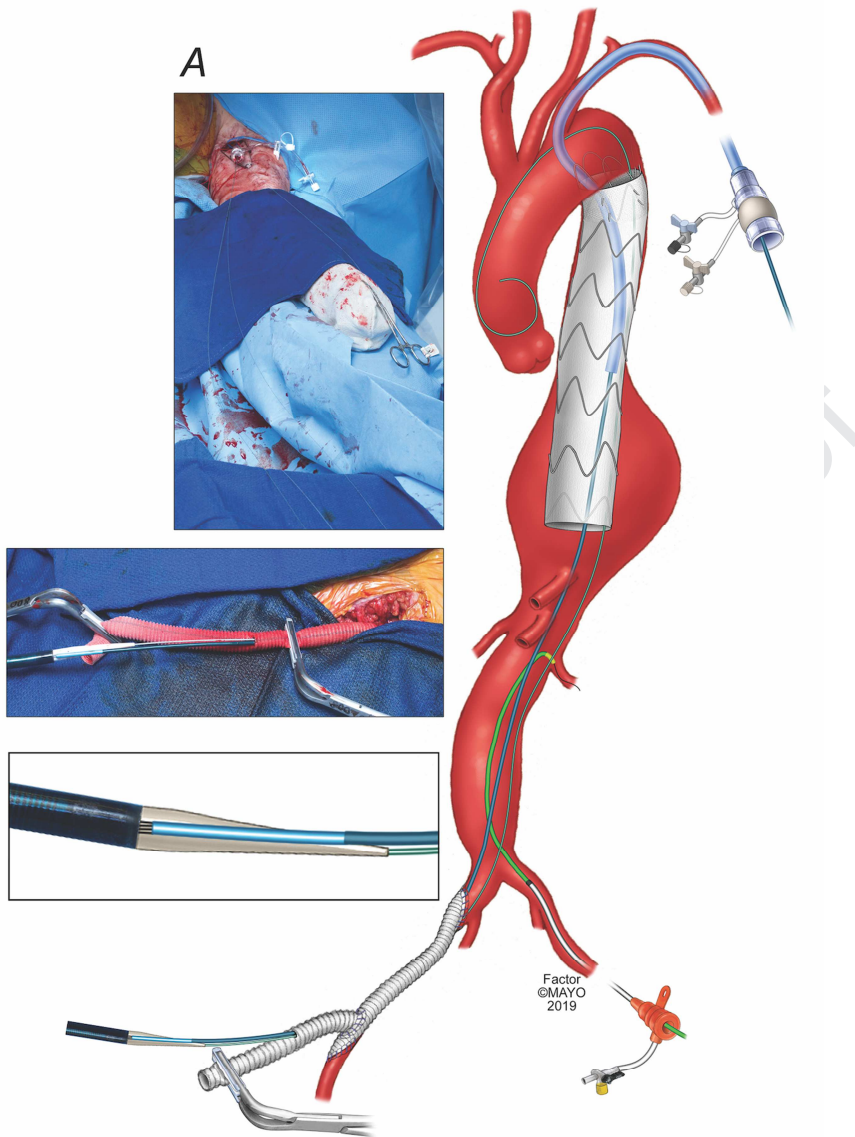
486

487

488

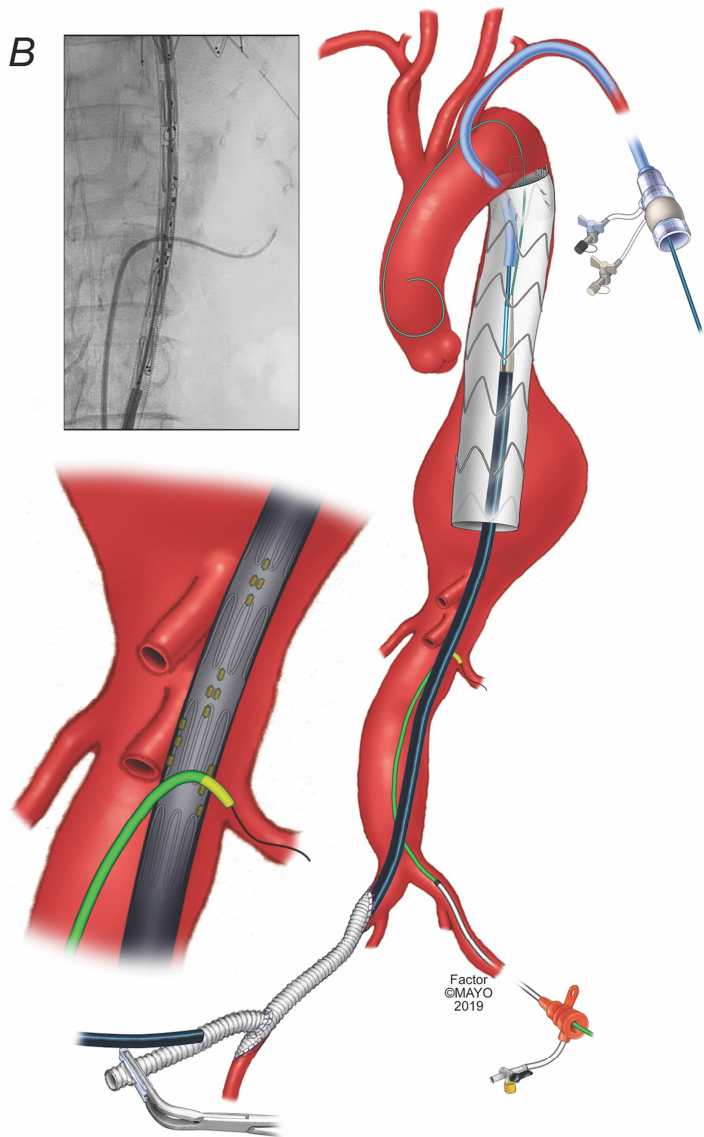
489

490 **Figure 4.** Through-and-through brachiofemoral access was established, then a 5Fr x  
491 90cm sheath was placed over this access. The preloaded wires were passed through the  
492 5-Fr through-and-through sheath (**A**). The modified stent graft was advanced along with  
493 the 5Fr brachiofemoral sheath. The stent graft was positioned under fusion imaging  
494 guidance, and was deployed in staggered fashion (**B**), allowing sequential branch  
495 catheterization using the preloaded wires (**C**). Following placement of branch bridging  
496 stents (**D-G**), and completion angiography showed patent branch stents and no endoleak  
497 (**H**). By permission of Mayo Foundation for Medical Education and Research. All rights  
498 reserved.



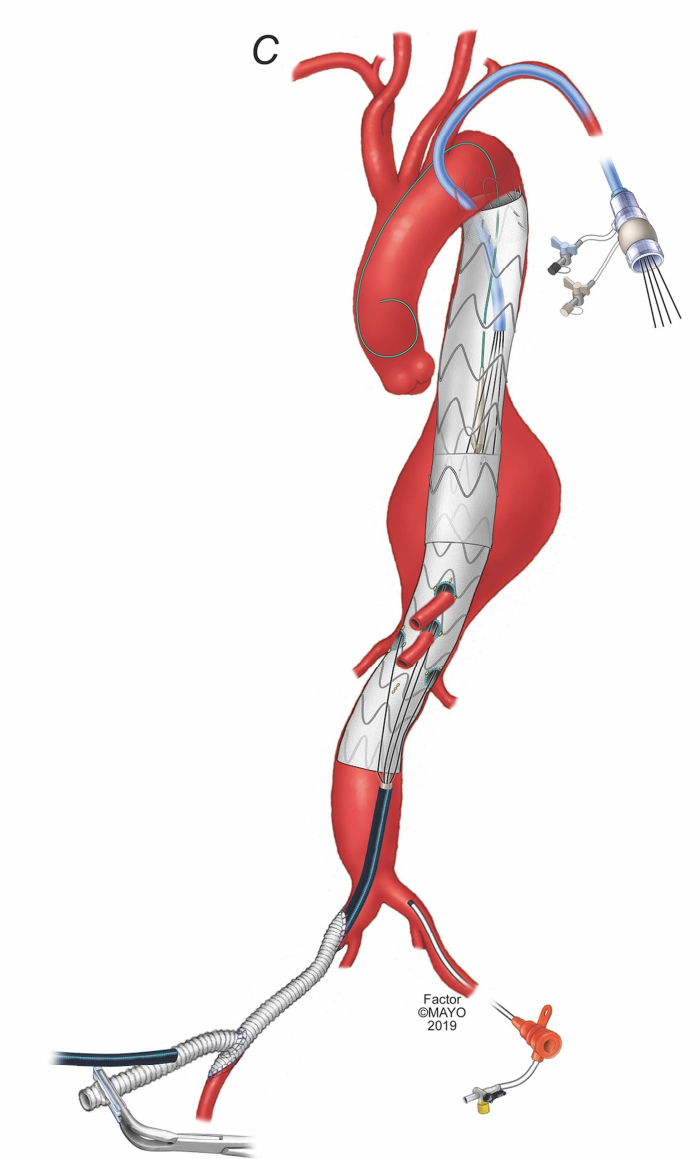
499

500



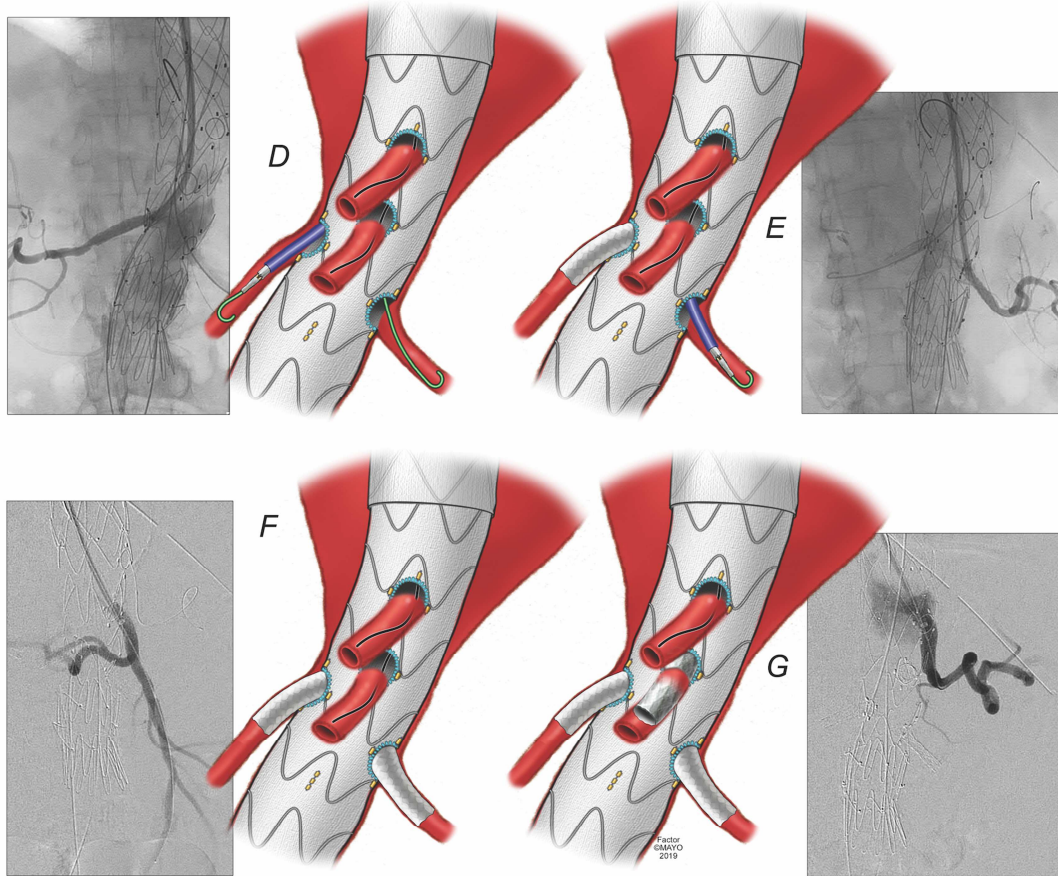
501

502



503

504







e-proof

506

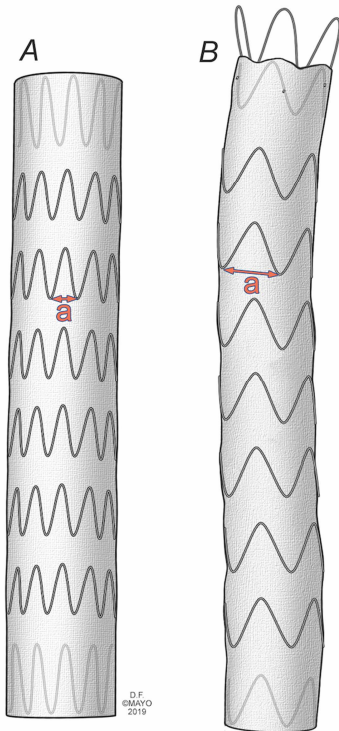
507

508

509

510

511 **Figure 5.** Distance apex to apex range (a) from 9.4mm ~ 11mm for Zenith TX2<sup>®</sup> thoracic  
512 stent graft (A), and 18.1mm ~ 18.8mm for Zenith Alpha<sup>™</sup> thoracic stent graft (B). By  
513 permission of Mayo Foundation for Medical Education and Research. All rights reserved.  
514



515

516

517

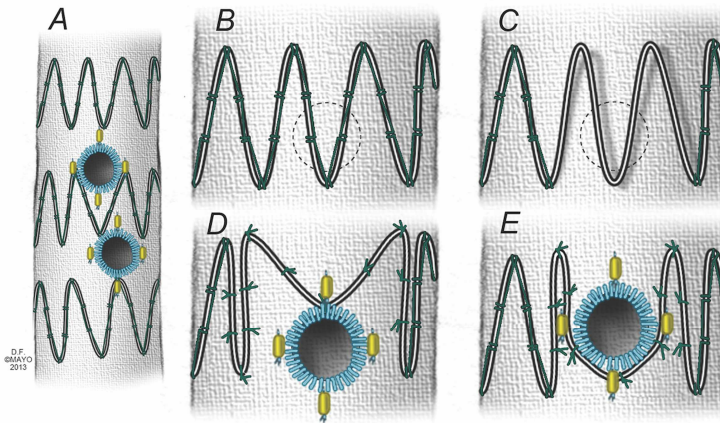
518

519

520

521

522 **Figure 6.** Stent relocation technique to accommodate fenestrations in Zenith TX2®  
523 thoracic stent graft requires bending and resuturing the Z stents (**D-E**). By permission of  
524 Mayo Foundation for Medical Education and Research. All rights reserved.



525

526

527

528

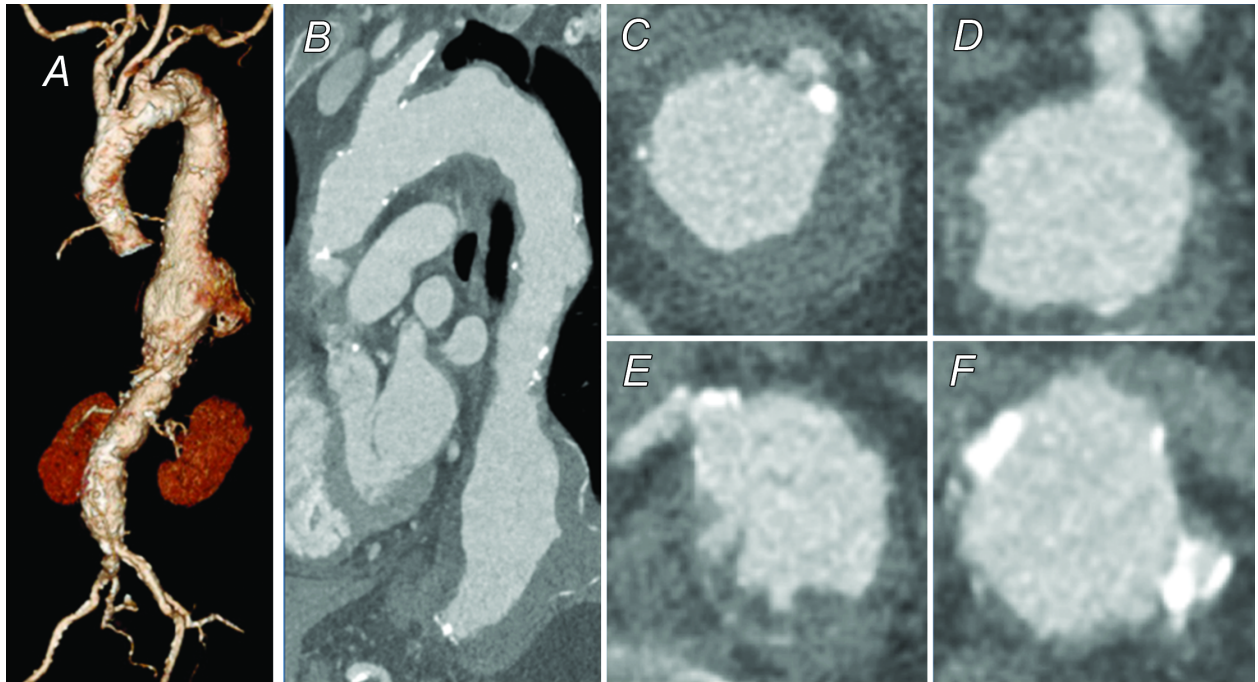
529

530 **Video.** Physician modified endograft (PMEG) technique using the low-profile Zenith

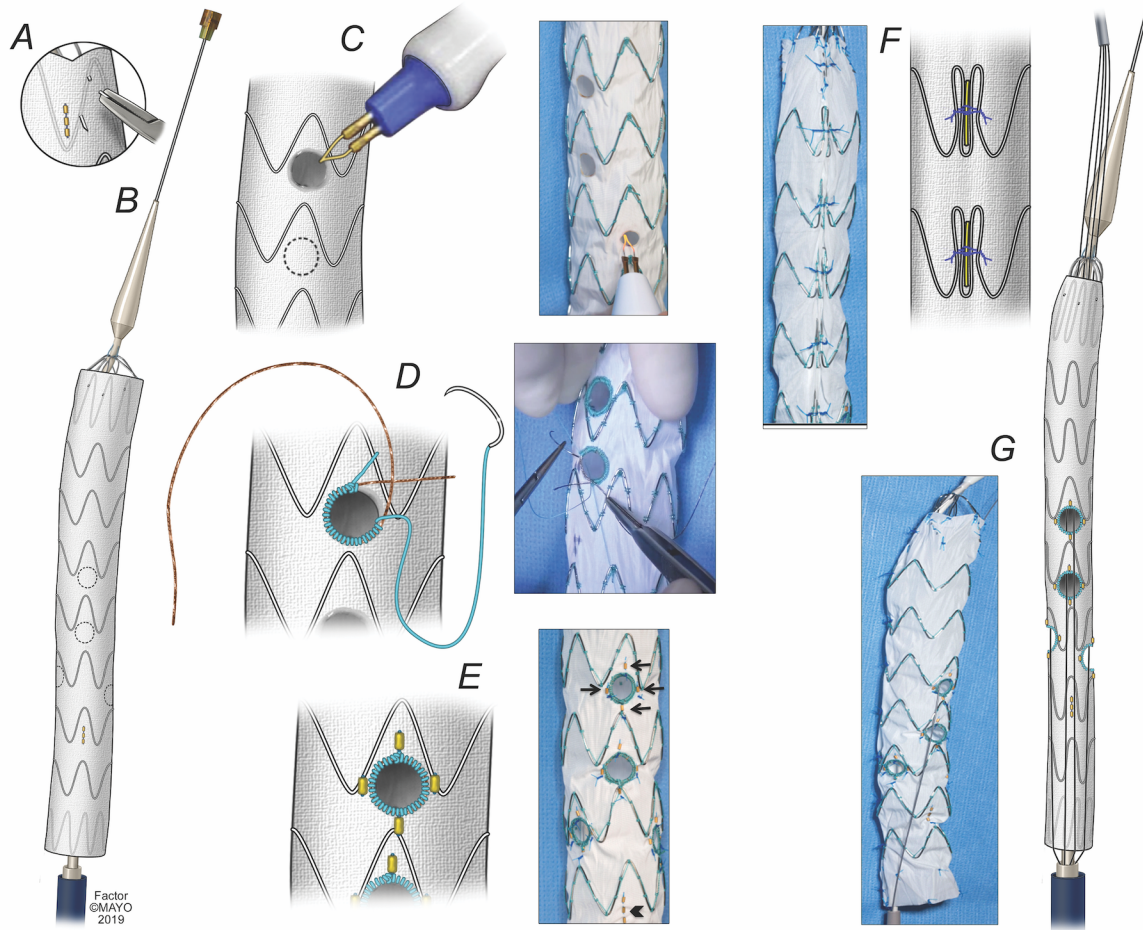
531 Alpha™ thoracic stent graft. By permission of Mayo Foundation for Medical Education

532 and Research. All rights reserved.

533

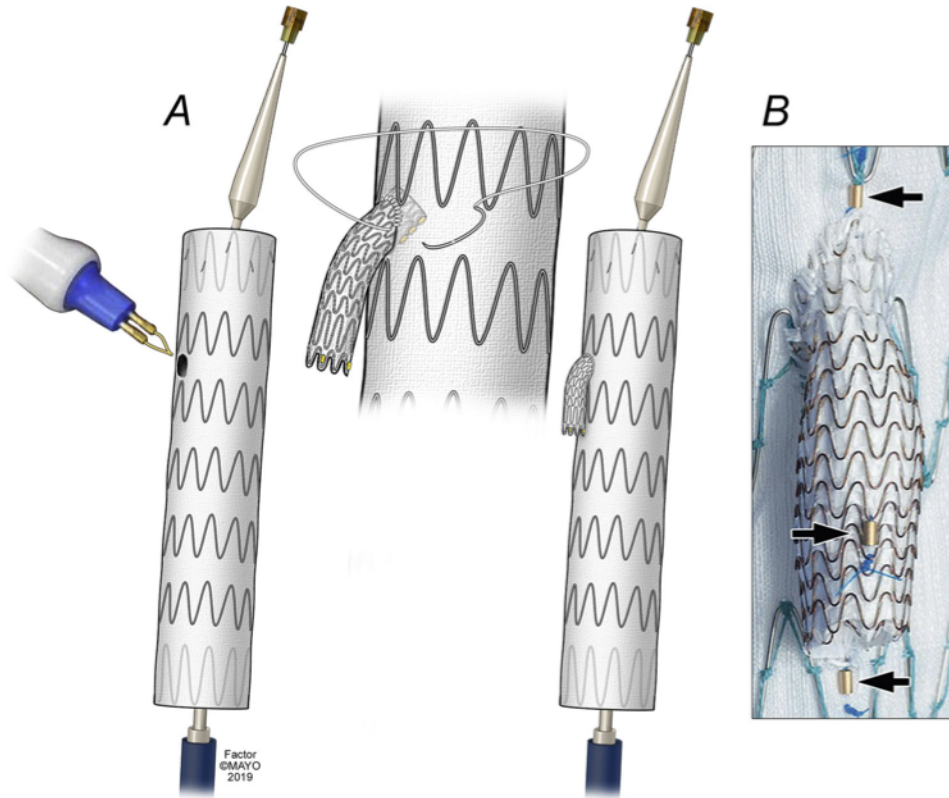


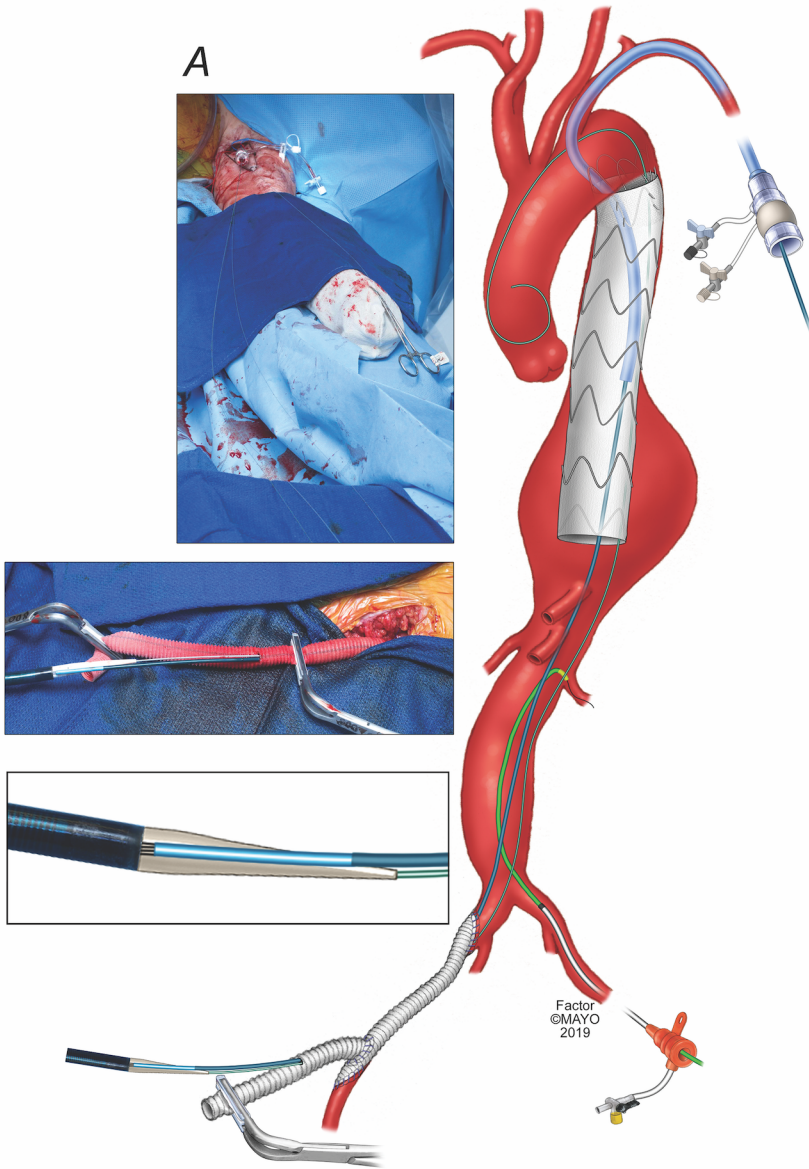
Journal Pre-proof



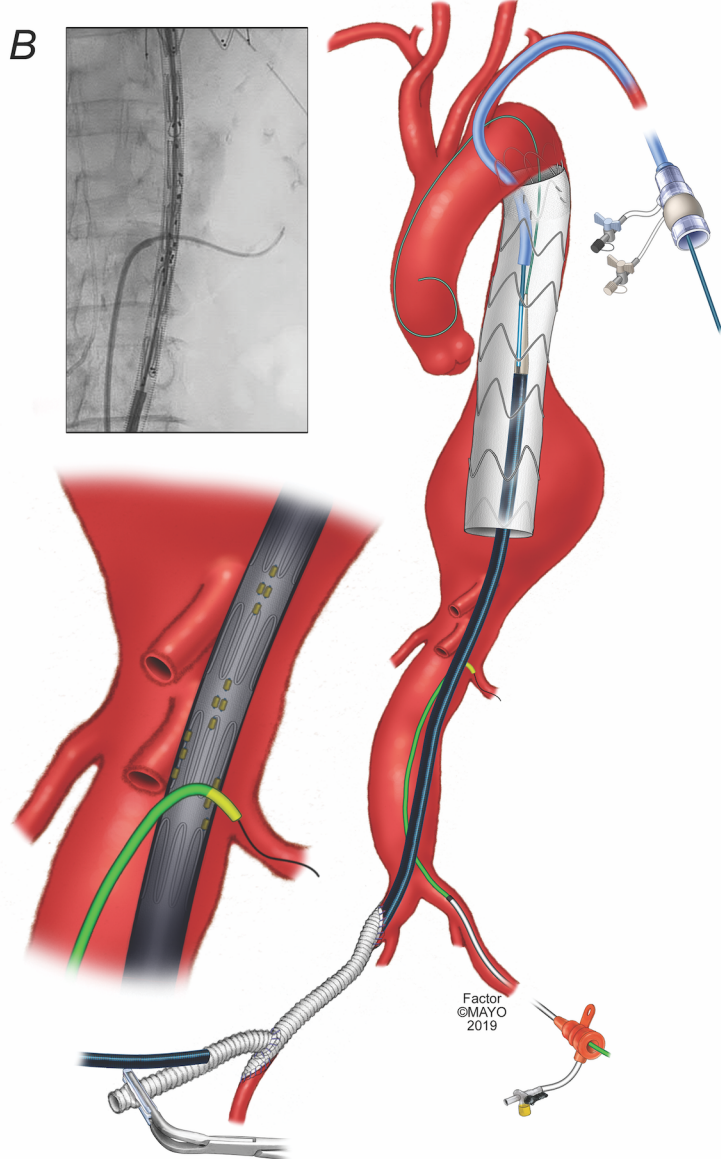
Journal







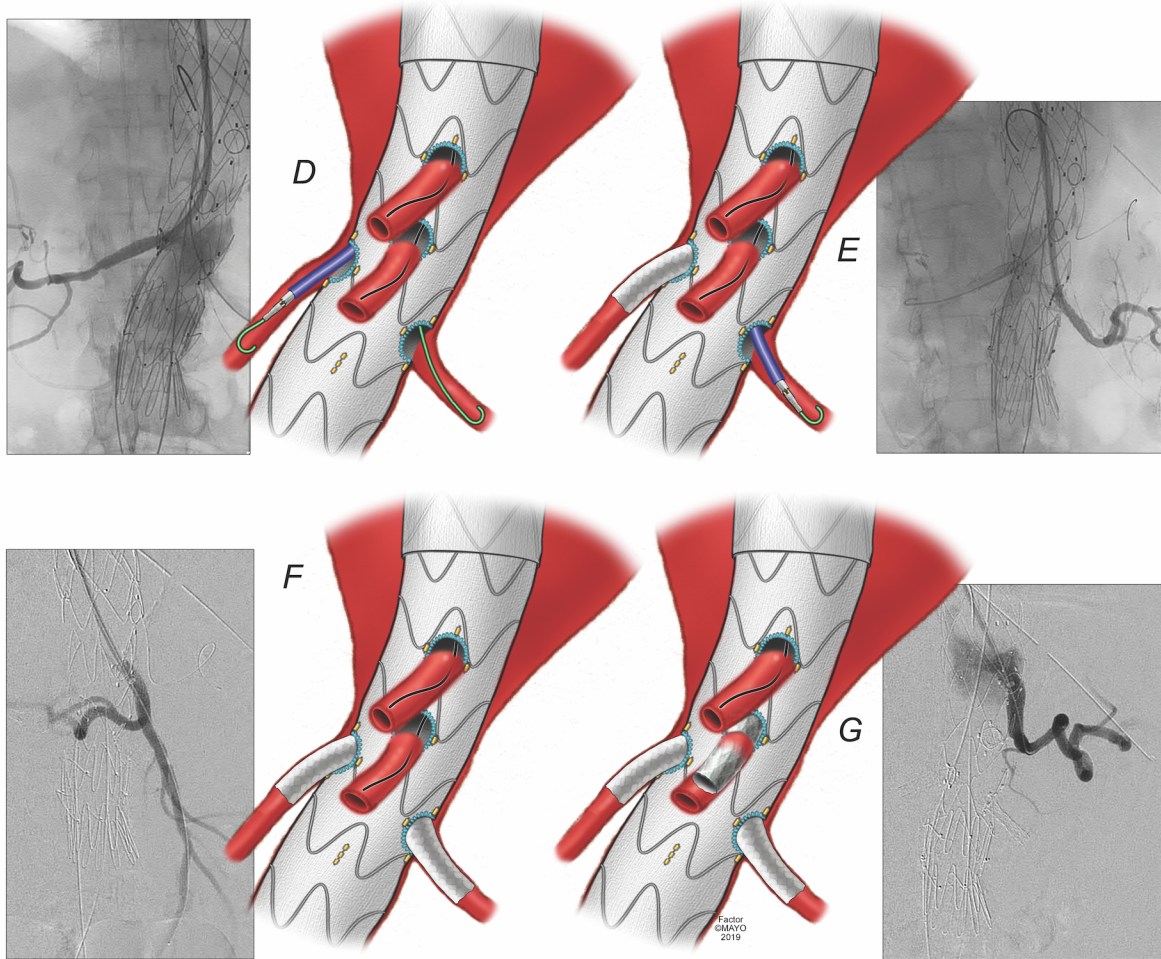




proof

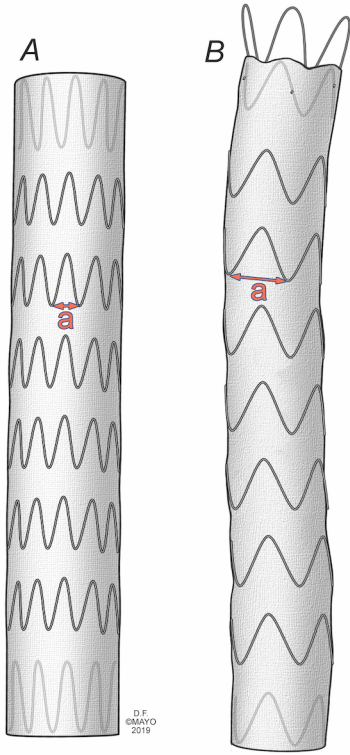


proof

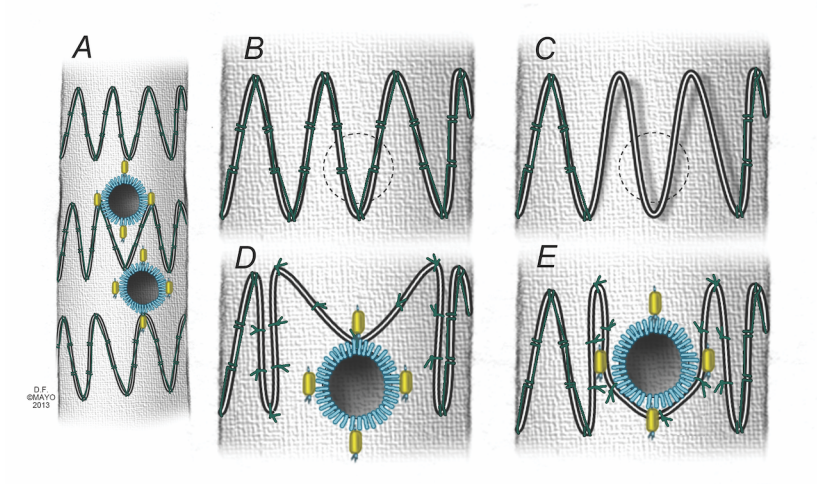




pre-proof



Journal Pre-proof



Journal Pre-proof